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INFORMATION DISCLOSURE STATEMENT

ATTORNEY DOCKET NO.: MEE-002

APPLICANTS: Miller et al.

SERIAL NO.: 09/780,142

FILING DATE: February 9, 2001

EXAMINER. Huynh, P.

GROUP: 1644

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EXAM. INIT.		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB CLASS	FILING DATE IF APPROPRIATE
<u> </u>	A1	5,707.986	1/13/98	Miller et al.	514	185	
	A2	5,733.876	3/31/98	O'Reilly et al.	514	12	
	A3	5,798.349	8/25/98	Levy et al.	514	185	
	14	5,854.205	12/29/98	O'Reilly et al.	512	2	
	A5	6,117.862	9/12/00	Margaron et al.	514	185	
	A6	6,180,402	1/30/01	Granville et al.	435	375	
	A7	6,225,303	5/1,01	Miller et al.	514	185	

## FOREIGN PATENT DOCUMENTS

EXAM. INIT.		DOCUMENT NUMBER	DATE	COUNTRY CODE	CLASS	SUB CLASS	FILING DATE	ABSTRACT ONLY	ENGLISH LANG (Y/N)
	B1 ,	WO 95 24930 A	09'21/95						
	B2 +	WO 97 33619 A	09 18/97						
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	B4 a	WO 98 25648 A	06 18/98						
	B5 🙃	WO 99 03503 A	01/28/99						
-	В6 е	WO 00 51638	09/08/00						
	B7	WO 01 51087 A	07/19/01						

## OTHER ART, JOURNAL ARTICLES, EXC.

EXAM. INIT.	OTHER DOCUMENTS: (Including Author, Title, Date, Relevant Pages, Place of Publication)				
	Cl	Adamis et al. "Inhibition of Vascular Endothelial Growth Factor Prevents Retinal Ischemia – Associated Iris Neovascularization in a Nonhuman Primate" (1996) Arch Ophthalmol, Vol. 114, pp. 66-71.			
	(2.	Ahmad et al "Photodynamic Therapy Results in Induction of WAFI/C1P1/P21 Leading to Cell Cycle Arrest and Apoptosis" (1998) Proc. Natl. Acad. Sci. USA. Vol. 95, pp: 6977 – 6982.			
	(3	Allen et al. "Photodynamic Therapy: Tumor Targeting with Adenoviral Proteins" (1999) Photochemistry and Photobiology, Vol.70, No. 4, pp. 512-523.			
	C4	Akhlynina et al. "In adin-mediated Intracellular Targeting Enhances the Photodynamic Activity of Chlorin e <sub>6</sub> " (1995) <u>Cancer Research</u> . Vol 55, pp1014-1019.			

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INFORMATION US 09/780,142	Sheet 2 of 5
C5 7	Arenberg et al. "Interferon-y-inducible 10 (IP-10) Is an Angiostatic Pactor That Inhibits Human Normall Cell Lung Canter (NSCLC) Tumorigenesis and Spontaneous Metastases" (1996) J. Exp. Med. Vol. 184, pp 981-992.
C6 5	Ball et al. "A Comparative Study of the Cellular Uptake and Photodynamic Efficacy of Three Novel Zinc Phthalocyanines of Differing Charge" (1999) Photochemistry and Photobiology. Vol. 69, No.3, pp. 390-39
	Berg, K. and Moan, J. "Lysosomes and Microtubules as Targets for Photochemotherapy of Cancer" (1997)  Photochemistry and Photobiology, Vol. 65, No. 3, pp. 403-409.
(8)	Biolo et al. "Effect of Photosensitizer Delivery System and Irradiation Parameters on the Efficiency of Photodynamic Therapy of B16 Pigmented Melanoma in Mice" (1996) <u>Photochemistry and Photobiology</u> , Vol. 63, No. 2, pp. 224-228.
(.8	Birchler M et al. "Selective Targeting and Photocoagulation of Ocular Angiogenesis Mediated by a Phagederived Human antibody Fragment" (1999) Nature Biotechnology, Vol. 17, No. 10, pp. 984-988.
C10	Brem et al. "The Combination of Antiangiogenic Agents to Inhibit Primary Tumor Growth and Metastasis" (1993) J. Ped. Surgery, Vol. 28, No. 10, pp. 1253-1257.
CH	Bressler, et al. 'Age-related Macular Degeneration" (1998) <u>Survey of Ophthalmology</u> , Vol. 32, No. 6, pp: 375-413.
C12	Brooks et al. "Integrin α <sub>ν</sub> β <sub>3</sub> Antagonists Promote Tumor Regression by Inducing Apoptosis of Angiogenic Blood Vessels" (1994) <u>Cell.</u> Vol. 79, pp. 1157-1164.
C13	Brooks et al. "Requirement of Vascular Integrin α <sub>ν</sub> β <sub>3</sub> for Angiogenesis" (1994) <u>Science</u> , Vol. 264, pp. 569-571.
C14	Ciardiello et al. "Antitumor Activity of Combined Blockade of Epidermal Growth Factor Receptor and Protein Kinase A" (1996) J. Natl. Can. Inst. Vol. 88, No. 23, pp. 1770-76.
C15	Ciulla T A et al. "Changing Therapeutic Paradigms for Exudative Age-related Macular Degeneration: Antiangiogenic Agents and Photodynamic Therapy" (1999) Expert Opinion on Investigational Drugs, Ashley Publications LTD, Vol. 8, No. 12, pp. 2173-2182.
C16	D'Amato et al "Thalidomide is an Inhibitor of Angiogenesis" (1994) Proc. Natl. Acad. Sci. USA, Vol. 91, pp: 4082-4085.
C17	Dimitroff C J et al. "Anti-Angiogenic Activity Of Selected Receptor Tyrosine Kinase Inhibitors, PD166285 And PD173074: Implications For Combination Treatment With Photodynamic Therapy" (1999)  Investigational New Drugs, Vol. 17, No. 2, pp: 121-35.
C18	Dobi et al. "A New Model of Experimental Choroidal Neovascularization in the Rat" (1989) Arch Ophthalmol, Vol. 107, pp. 264-269.
C19	Dougherty et al. "Photodynamic Therapy" (1998) J. Natl. Can. Inst., Vol. 90, No. 12, pp. 889-905.
C20	Ellerby et al. "Anti-cancer activity of targeted pro-apoptotic peptides" (1999) Nature Medicine, Vol. 5, No. 9, pp: 1032-1038.
C21	Ferrario et al. "Antiangiogenic Treatment Enhances Photodynamic Therapy Responsiveness in a Mouse Mammary Carcinoma" (2000) Cancer Research, Vol. 60, No. 15, pp. 4066-4069.
C22	Fingar et al. "Analysis of Acute Vascular Damage After Photodynamic Therapy Using Benzoporphyrin Derivative (BPD)" (1999) British Journal of Cancer, Vol. 79, pp. 1702-1708.
C23	Folkman, J. "Angiogenesis in Cancer, Vascular, Rheumatoid and Other Disease" (1995) <u>Nature Medicine</u> , Vol. 1, pp. 27-31.
(24	Friedlander et al. "Definition of Two Angiogenic Pathways by Distinct α <sub>0</sub> Integrins"(1995) <u>Science</u> . Vol. 270, pp. 1500-1502.

	Sheet 3 of 6
*	Friedlander et al. "Involvement of Integrins α <sub>0</sub> β <sub>3</sub> and α <sub>0</sub> β <sub>5</sub> in Ocular Neovascular Diseases" (1996) Pro. Natl. Acad. Sci. USA. Vol. 93. pp. 9764-9769.
C26	Granville et al. "Overexpression of Bcl-X <sub>L</sub> Prevents Caspase-3-mediated Activation of DNA Fragmentation Factor (DFF) Produced by Treatment with the Photochemotherapeutic Agent BPD-MA" (1998) <u>FEBS</u> Letters, Vol. 422. pp: 151-154.  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31  Granville et al. "Granville cytochrome c Release, Activation of Caspases 3, 6,
C27	Granville et al. "Rapid cytochrome c Release, Activation of Caspases 3, 6, 7 and 8 Followed by Bap31 Cleavage in HeLa cells Treated with Photodynamic Therapy" (1998) FEBS Letters, Vol. 437, pp. 5-10.
C28	Oupta et al. "Involvement of Nitric Oxide During Phthalocyanine (Pc4) Photodynamic Therepy-Mediated Aportosis" (1998) Cancer Research, Vol. 58, pp. 1785-1788.
C29	Guyer et al. "Subfoveal Choroidal Neovascular Membranes in Age-Related Macular Degeneration" (1986) <u>Arch Ophhalmol</u> , Vol. 104 pp; 702-705.
 C30	Hakem et al. Differential Requirement for Caspase 9 in Apoptotic Pathways in Vivo" (1998) Cell, Vol. 94, pp: 339-352.
C31	Hanahan, D. and Folkman, J. "Patterns and Emerging Mechanisms of the Angiogenic Switch during Tumorigenesis" (1996). Cell, Vol. 86, pp. 353-364.
C32	He et al. "Photodynamic Therapy with Photofrin II Induces Programmed Cell Death in Carcinoma Cell Lines" (1994) Photochemistry and Photobiology, Vol. 59, No. 4, pp; 468-473.
C33	He et al. "The Induction of Partial Resistance to Photodynamic Therapy by the Protooncogene <i>BCL-2</i> " (1996) <u>Photochemistry and Photobiology</u> , Vol. 64, No. 5, pp. 845-852.
C34	He et al. 'Variation in Photodynamic Efficacy during the Cellular Uptake of Two Phthalocyanine Photosensitizers' (1998) Photochemistry and Photobiology. Vol. 67, No. 6, pp. 720-728.
C35	Hockenbery et al. "Bel-2 Functions in an Antioxidant Pathway to Prevent Apoptosis" (1993) Cell, Vol. 75, pp. 241-251.
C36	Hunt David W.C. et al. "Impact of PDT and Fas-mediated Apoposis on Cells of the System" (1998)  Photochemistry and Photobiology, Vol. 67, Spec. Issue, pp. 69S.
C37	2 (Ama Balatad Manufar Degeneration) Chapter 17, pp. 297-307.
C38	Husain, et al. 'Intravenous Infusion of Liposomal Benzoporphyrin Derivative for Photodynamic Therapy of Experimental Choroidal Neovascularization' (1996) <u>Arch Ophthalmol</u> , Vol. 114, pp. 978-985.
C39	Hyman et al. 'Senile Macular Degeneration: A Case-Control Study (1983) American Journal of Epidemiology, Vol. 118, No. 2., pp. 213-227.
C40	Ingher et al. "Synthetic Analogues Of Fumagillin That Inhibit Angiogenesis And Suppress Tumour Growth" (1990) Nature, Vol. 348, pp. 555-557.
C41	Kane et al. "Bcl-2 Inhibition of Neural Death: Decreased Generation of Reactive Oxygen Species" (1993)  Science, Vol. 262, No. 19, pp. 1274-1277.
C42	Kerr, et al. "Apoptosis: A Basic Biological Phenomenon with Wide-Ranging Implications in Tissue Kinetics" (1972) Br. J. Caneer, Vol. 26, pp. 239-257.
 C43	Klessel et al. Biodistribution of Photosensitizing Agents" (1993) Int. J. Biochem, Vol. 25, No. 10, pp. 1377-83.
(*44	V rouge 1 1 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2
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	C46	Kramer, et al. "Liposomal Benzoporphyrin Derivative Verteporfin Photodynamic Therapy" (1996)  Ophthalmology Vol. 103, No. 3, pp. 427-438.  Kroemer, G. "The Proto-oncogene Bel-2 and Its Role in Regulating Apoptosis" (1997) Nature Medicine, Vol. 3, No. 6, pp. 614-620.  Lucas et al. "Multiple Forms of Angiostatin Induce Apotosis in Endothelial Cells" (1998) Blood, Vol. 92.
	C47	Kroemer, G. "The Proto-oncogene Bel-2 and Its Role in Regulating Apoptosis" (1997) Nature Medicine, Vol. 3. No. 6, pp. 614-620.
	C48	Lucas et al. "Multiple Forms of Angiostatin Induce Apotosis in Endothelial Cells" (1998) <u>Blood</u> , Vol. 92. No. 12, pp: 4730-4741.
	C49	Luo et al. "The Role of Mitochondrial Photodamage in PDT-induced Apoptosis" (1998) Chemical Abstracts Vol. 129, No. 13 & Proc. Spie-Int. Soc. Opit. Eng. Vol. 3247, pp. 112-117.
	C50	Manceri et al. "Combined effects of Angiostatin and Ionizing Radiation in Antitumour Therapy" (1998)  Nature Vol. 394, pp. 287-291.
	C51	Miller et al. "Alternative Treatments of Neovascular Age-Related Macular Degeneration" (1999) Chapter 139, pp 2014-2027.
_	C52	Miller JW et al. Photodynamic Therapy of Experimental Chorodial Neovascularization Using Lipoprotein-deliver Benzoporphyrin' (1995) <u>Archives of Opthalmology</u> , Vol. 113, No. 6, pp. 810-818.
	C53	Mori et al. "Photodynamic Therapy for Experimental Tumors Using ATX-S10(Na), a Hydrophilic Chlorin Photosensitizer, and Diode Laser." (2000) <u>Ipn. J. Cancer.</u> Vol. 91, No. 7, pp: 753-9.
	C54	Moser et al. "Angiostatin binds ATP Synthase on the Surface of Human Endothelial Cells" (1999) Proc. Natl Acad. Sci USA, Vol. 96, pp; 2811-2816.
<u></u>	C55	Mosmann, T. "Rapid Colorimetric Assay for Cellular Growth and Survival: Application to Proliferation and Cytotoxicity Assays" (1983) Journal of Immunological Methods, Vol. 65, pp. 55-63.
	C56	1200 1200
<u></u>	C57	Nicholson, D. and Thornberry, N. "Capases: Killer Proteases" (1997) TIBS, Vol. 22, pp. 299-306.
	C58	Obana et al. "Selective Occlusion of Choroidal Neovascularization by Photodynamic Therapy with a Water-Soluble Photosensitizer, ATX-S10" (1999) <u>Lasers Surg. Med.</u> , Vol. 24, No. 3, pp. 209-222.
	C59	Obana et al. "Selective Photodynamic Effects of the New Photosensitizer ATX-S10(Na) on Choroidal Neovascularization in Monkeys" (2000) <u>Arch. Ophthalmol.</u> Vol. 118, No. 5, pp. 650-658.
	C60	20 1 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2
	C61	Oleinick et al "The Photobiology of Photodynamic Therapy: Cellular Targets and Mechanisms" (1998) Radiation Research, 150 (Suppl.), pp. S146-S156.
	C62	O'Reilly et al. "Angiostatin Induces And Sustains Dormancy Of Human Primary Tumors In Mice" (1996)  Nature Medicine, Vol. 2, No. 6, pp. 689-692.
	C63	O'Reilly et al. "Angiostatin: A Novel Angiogenesis Inhibitor That Mediates the Superession of Metastases by a Lewis Lung Carcinoma" (1994) Cell, pp. 315-328.
	C64	
	C65	Pidgeon & Colles et al. "Recent Developments In Tunable Lasers For Spectroscopy" (1979) Nature. Vol. 279, pp: 377-381.
	C60	Pollack et al. "Repair of Retinal Pigment Epithelium and Choriocapillaries After Laser Photocoagulation: Correlations Between Scanning Electron, Transmission Electron and Light Microscopy" (1997) Opthal. Res Vol. 29, pp: 393-404.

		Sheet 5 of C
	C67	Reddi, E. "Role of Delivery Vehicles for Photographizers in the Photodynamic Therapy of Tumors" (1997)  J. Photochem & Photobiol. Vol 37, pp 189-195.
	C68	Renno et al. "Photodynamic Therapy Using Lu-Tex Induces Apoptosis In Vitro, and its Effect is Potentiated by Angiostatin in Retinal Capillary Endothelial Cells" (2000) Inves. Ophthal. & Vis. Sci. Vol. 41. No. 12, pp: 3963-71.
	(20)	Ryan "Subretinal Neovascularization: Natural History of an Experimental Model" (1982) <u>Arch Ophthalmol,</u> Vol. 100, pp. 1804-1809.
-	C70	Schmidt-Erfurth, U. et al. "Photodynamic therapy in Ocular Vascular Disease" (1998) <u>Laser Phys.</u> , Vol. 8, No. 1, pp. 191-198.
	C71	Senior, K."Ageing eyes retain their mystery." (1999) The Lancet Vol. 353, pp. 818.
	C72	Separovic et al "Association of Ceramide Accumulation with Photodynamic Treatment-Induced Cell Death" (1998) Photochemistry and Photobiology, Vol. 68, No. 1, pp. 101-109.
	C73	Sharma A. et al. "Combination of Photodynamic Therapy and Angiogenic Therapy for the Treatment of Human Breast Cancer in Nude Athymic Myce" (1996) Proceedings of the Annual Meeting of the American Association for Cancer Research, US, Phildelphia, AACR, Vol. Meeting 87, pp. 291.
	C74	riginesii ripitiiettiini-Detrived Litetor (2000) = 1
	C75	Tolentino et al. "Vascular Endothelial Growth Factor Is Sufficient to Produce Iris Neovascularization and Neovascular Glaucoma in a Nonhuman Primate" (1996) <u>Arch Ophthalmol</u> , Vol. 114, pp. 964-969.
	C76	Varnes et al. "Photodynamic Therapy Induced Apoptosis in Lymphoma Cells: Translocation of Cytochrome e Causes Inhibition of Respiration as Well as Caspase Activation" (1999) <u>Biochem. &amp; Biophys. Res. Com.</u> , Vol. 255, pp. 673-679.
	C77	Veis, et al. "Be1-2-Deficient Mice Demonstrate Fulminant Lymphoid Apoptosis, Polycystic Kidneys, and Hypopigmented Hair" (1993) Cell, Vol. 75, pp. 229-240.
	C78	Virgill et al. "Bc1-2 Overexpression in the HaCaT Cell Line Is Associated with a Different Membrane Fatty Acid Composition and Sensitivity To Oxidative Stress" (1998) Free Radical Biology & Medicine, Vol. 24, No. 1, pp. 93-101.
	C79	Voest et al. "Inhibition of Angiogenesis In Vivo by Interleukin 12" (1995) J. Nat. Can. Inst., Vol. 87, No. 8 pp: 581-586.
	C80	Antibodies: A New Possibility for Use in Photodynamic Therapy" (2000) Int. J. Cancer, Vol. 88, pp. 108-114.
	C81	Weishaupt et al. "Identification of Singlet Oxygen as the Cytotoxic Agent in Photo-inactivation of a Murine Tumor" (1976) Can. Res., Vol. 36, pp. 2326-2329.
	C82	Cultivity Different Proliferative States" (1990) Int. J. Radiat. Biol., Vol. 58, No. 1, pp. 145-156.
	C83	Woodburn et al. "Localization and Efficacy Analysis of the Phototherapeutic Lutetium Texaphyrin (PCl-0123) in the Murine EMT6 Sarcoma Model" (1997) Photochemistry and Photobiology, Vol. 65, No. 3, pp 410-415.
	(*8-	Phytochemistry and Photobiology, Vol. 63, No. 6, pp. 892-897.
	C8.	Zhang et al. "Antisense Bc1-2 Retrovirus Vector Increases the Sensitivity of a Human Gastric Adenocarcinoma Cell Line to Photodynamic Therapy" (1999) <u>Photochemistry and Photobiology</u> , Vol. 69, No. 5, pp. 582-586.

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Sheet 6 of 6

C86 International Preliminary Examination Report for PCT/US01/04231 dated May 12, 2001 (5 pages):

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